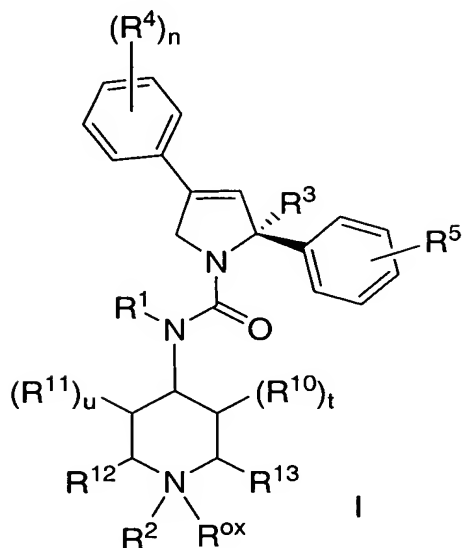


In the claims:

1. (Original) A compound of Formula I:



or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

- a is 0 or 1;
b is 0 or 1;
m is 0, 1, or 2;
n is 0, 1, 2 or 3;
r is 0 or 1;
s is 0 or 1;
t is 0, 1 or 2;
u is 0 or 1;

R¹ and R² are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷;

R³ is selected from:

- 1) hydrogen;

- 2) C₁-C₁₀ alkyl;
- 3) C₁-C₁₀ alkyl-O-R^d,
- 4) C₂-C₁₀ alkenyl-O-R^d,
- 5) C₂-C₁₀ alkynyl-O-R^d,
- 6) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-O-R^d,
- 7) C₁-C₁₀ alkyl-(C=O)_b-NR^cR^{c'},
- 8) C₂-C₁₀ alkenyl-(C=O)_b-NR^cR^{c'},
- 9) C₂-C₁₀ alkynyl-(C=O)_b-NR^cR^{c'},
- 10) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-(C=O)_b-NR^cR^{c'},
- 11) C₁-C₁₀ alkyl-S(O)_m-R^d,
- 12) C₂-C₁₀ alkenyl-S(O)_m-R^d,
- 13) C₂-C₁₀ alkynyl-S(O)_m-R^d,
- 14) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-S(O)_m-R^d,

said alkyl, alkenyl, alkynyl and cycloalkyl are optionally substituted with one or more substituents selected from R⁶;

R⁴ is independently selected from:

- 1) (C=O)_aO_bC₁-C₁₀ alkyl,
- 2) (C=O)_aO_baryl,
- 3) CO₂H,
- 4) halo,
- 5) CN,
- 6) OH,
- 7) O_bC₁-C₆ perfluoroalkyl,
- 8) O_a(C=O)_bNR⁸R⁹,
- 9) S(O)_mR^a,
- 10) S(O)₂NR⁸R⁹, and
- 11) -OPO(OH)₂;

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R⁷;

R⁵ is selected from:

- 1) hydrogen;
- 2) (C=O)_aO_bC₁-C₁₀ alkyl,

- 3) $(\text{C}=\text{O})_a\text{O}_b\text{aryl}$,
- 4) CO_2H ,
- 5) halo,
- 6) CN ,
- 7) OH ,
- 8) $\text{O}_b\text{C}_1\text{-C}_6$ perfluoroalkyl,
- 9) $\text{O}_a(\text{C}=\text{O})_b\text{NR}^8\text{R}^9$,
- 10) $\text{S}(\text{O})_m\text{R}^a$,
- 11) $\text{S}(\text{O})_2\text{NR}^8\text{R}^9$, and
- 12) $-\text{OPO}(\text{OH})_2$;

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R^7 ;

R^6 is independently selected from:

- 1) $(\text{C}=\text{O})_a\text{O}_b\text{C}_1\text{-C}_{10}$ alkyl,
- 2) $(\text{C}=\text{O})_a\text{O}_b\text{aryl}$,
- 3) $\text{C}_2\text{-C}_{10}$ alkenyl,
- 4) $\text{C}_2\text{-C}_{10}$ alkynyl,
- 5) $(\text{C}=\text{O})_a\text{O}_b$ heterocyclyl,
- 6) CO_2H ,
- 7) halo,
- 8) CN ,
- 9) OH ,
- 10) $\text{O}_b\text{C}_1\text{-C}_6$ perfluoroalkyl,
- 11) $\text{O}_a(\text{C}=\text{O})_b\text{NR}^8\text{R}^9$,
- 12) $\text{S}(\text{O})_m\text{R}^a$,
- 13) $\text{S}(\text{O})_2\text{NR}^8\text{R}^9$,
- 14) oxo,
- 15) CHO ,
- 16) $(\text{N}=\text{O})\text{R}^8\text{R}^9$,
- 17) $(\text{C}=\text{O})_a\text{O}_b\text{C}_3\text{-C}_8$ cycloalkyl, and
- 18) $-\text{OPO}(\text{OH})_2$;

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R^7 ;

R⁷ is selected from:

- 1) (C=O)_rO_s(C₁-C₁₀)alkyl,
- 2) O_r(C₁-C₃)perfluoroalkyl,
- 3) oxo,
- 4) OH,
- 5) halo,
- 6) CN,
- 7) (C₂-C₁₀)alkenyl,
- 8) (C₂-C₁₀)alkynyl,
- 9) (C=O)_rO_s(C₃-C₆)cycloalkyl,
- 10) (C=O)_rO_s(C₀-C₆)alkylene-aryl,
- 11) (C=O)_rO_s(C₀-C₆)alkylene-heterocyclyl,
- 12) (C=O)_rO_s(C₀-C₆)alkylene-N(R^b)₂,
- 13) C(O)R^a,
- 14) (C₀-C₆)alkylene-CO₂R^a,
- 15) C(O)H,
- 16) (C₀-C₆)alkylene-CO₂H, and
- 17) (C=O)_rN(R^b)₂,
- 18) S(O)_mR^a,
- 19) S(O)₂N(R^b)₂; and
- 20) -OPO(OH)₂;

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, alkylene and heterocyclyl is optionally substituted with up to three substituents selected from R^b, OH, (C₁-C₆)alkoxy, halogen, CO₂H, CN, O(C=O)C₁-C₆ alkyl, oxo, NO₂ and N(R^b)₂;

R⁸ and R⁹ are independently selected from:

- 1) H,
- 2) (C=O)O_bC₁-C₁₀ alkyl,
- 3) (C=O)O_bC₃-C₈ cycloalkyl,
- 4) (C=O)O_baryl,
- 5) (C=O)O_bheterocyclyl,
- 6) C₁-C₁₀ alkyl,
- 7) aryl,
- 8) C₂-C₁₀ alkenyl,

- 9) C₂-C₁₀ alkynyl,
- 10) heterocyclyl,
- 11) C₃-C₈ cycloalkyl,
- 12) SO₂R^a, and
- 13) (C=O)NR^b₂,

said alkyl, cycloalkyl, aryl, heterocyclyl, alkenyl, and alkynyl is optionally substituted with one, two or three substituents selected from R⁷, or

R⁸ and R⁹ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R¹⁰ and R¹¹ are independently selected from: F and -CH₂F;

R¹² and R¹³ are independently selected from: H and -CH₂F;

R^{ox} is absent or is oxo;

R^a is independently selected from: (C₁-C₆)alkyl, (C₃-C₆)cycloalkyl, aryl, or heterocyclyl, optionally substituted with one, two or three substituents selected from R⁷;

R^b is independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl, (C₃-C₆)cycloalkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^e ' or S(O)₂R^a, optionally substituted with one, two or three substituents selected from R⁷;

R^c and R^c ' are independently selected from: H, (C₁-C₆)alkyl, aryl, NH₂, OH, OR^a, -(C₁-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^e ', S(O)₂R^a and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷; or

R^c and R^c ' can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the

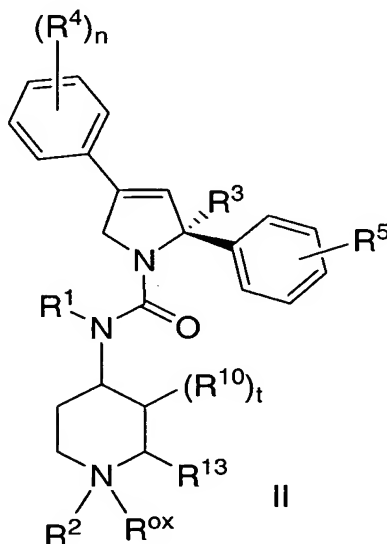
nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^d is selected from: H, (C₁-C₆)alkyl, -(C₂-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷;

R^e and R^{e'} are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷; or

R^e and R^{e'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷.

2. (Original) The compound according to Claim 1 of Formula II:



or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

a is 0 or 1;

b is 0 or 1;
m is 0, 1, or 2;
n is 0, 1, 2 or 3;
r is 0 or 1;
s is 0 or 1;
t is 0 or 1;

R¹ and R² are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷;

R³ is selected from:

- 1) hydrogen;
- 2) C₁-C₁₀ alkyl;
- 3) C₁-C₁₀ alkyl-O-R^d,
- 4) C₂-C₁₀ alkenyl-O-R^d,
- 5) C₂-C₁₀ alkynyl-O-R^d,
- 6) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-O-R^d,
- 7) C₁-C₁₀ alkyl-(C=O)_b-NR^cR^{c'},
- 8) C₂-C₁₀ alkenyl-(C=O)_b-NR^cR^{c'},
- 9) C₂-C₁₀ alkynyl-(C=O)_b-NR^cR^{c'},
- 10) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-(C=O)_b-NR^cR^{c'},
- 11) C₁-C₁₀ alkyl-S(O)_m-R^d,
- 12) C₂-C₁₀ alkenyl-S(O)_m-R^d,
- 13) C₂-C₁₀ alkynyl-S(O)_m-R^d,
- 14) (C₁-C₆-alkylene)_nC₃-C₈ cycloalkyl-S(O)_m-R^d,

said alkyl, alkenyl, alkynyl and cycloalkyl are optionally substituted with one or more substituents selected from R⁶;

R⁴ is independently selected from:

- 1) (C=O)_aO_bC₁-C₁₀ alkyl,
- 2) (C=O)_aO_baryl,
- 3) CO₂H,
- 4) halo,
- 5) CN,
- 6) OH,

- 7) $O_bC_1-C_6$ perfluoroalkyl,
- 8) $O_a(C=O)_bNR^8R^9$,
- 9) $S(O)_mR^a$,
- 10) $S(O)_2NR^8R^9$,

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R^7 ;

R^5 is selected from:

- 1) hydrogen;
- 2) $(C=O)_aO_bC_1-C_{10}$ alkyl,
- 3) $(C=O)_aO_b$ aryl,
- 4) CO_2H ,
- 5) halo,
- 6) CN ,
- 7) OH ,
- 8) $O_bC_1-C_6$ perfluoroalkyl,
- 9) $O_a(C=O)_bNR^8R^9$,
- 10) $S(O)_mR^a$,
- 11) $S(O)_2NR^8R^9$, and
- 12) $-OPO(OH)_2$;

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R^7 ;

R^6 is independently selected from:

- 1) $(C=O)_aO_bC_1-C_{10}$ alkyl,
- 2) $(C=O)_aO_b$ aryl,
- 3) C_2-C_{10} alkenyl,
- 4) C_2-C_{10} alkynyl,
- 5) $(C=O)_aO_b$ heterocyclyl,
- 6) CO_2H ,
- 7) halo,
- 8) CN ,
- 9) OH ,
- 10) $O_bC_1-C_6$ perfluoroalkyl,

- 11) $O_a(C=O)_bNR^8R^9$,
- 12) $S(O)_mR^a$,
- 13) $S(O)_2NR^8R^9$,
- 14) oxo,
- 15) CHO,
- 16) $(N=O)R^8R^9$,
- 17) $(C=O)_aO_bC_3-C_8$ cycloalkyl, and
- 18) $-OPO(OH)_2$;

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one, two or three substituents selected from R^7 ;

R^7 is selected from:

- 1) $(C=O)_rO_s(C_1-C_{10})$ alkyl,
- 2) $O_r(C_1-C_3)$ perfluoroalkyl,
- 3) oxo,
- 4) OH,
- 5) halo,
- 6) CN,
- 7) (C_2-C_{10}) alkenyl,
- 8) (C_2-C_{10}) alkynyl,
- 9) $(C=O)_rO_s(C_3-C_6)$ cycloalkyl,
- 10) $(C=O)_rO_s(C_0-C_6)$ alkylene-aryl,
- 11) $(C=O)_rO_s(C_0-C_6)$ alkylene-heterocyclyl,
- 12) $(C=O)_rO_s(C_0-C_6)$ alkylene- $N(R^b)_2$,
- 13) $C(O)R^a$,
- 14) (C_0-C_6) alkylene- CO_2R^a ,
- 15) $C(O)H$,
- 16) (C_0-C_6) alkylene- CO_2H ,
- 17) $C(O)N(R^b)_2$,
- 18) $S(O)_mR^a$,
- 19) $S(O)_2N(R^b)_2$; and
- 20) $-OPO(OH)_2$;

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, alkylene and heterocyclyl is optionally substituted with up to three substituents selected from R^b , OH, (C_1-C_6) alkoxy, halogen, CO_2H , CN, $O(C=O)C_1-C_6$ alkyl, oxo, NO_2 and $N(R^b)_2$;

R^8 and R^9 are independently selected from:

- 1) H,
- 2) $(C=O)O_bC_1-C_{10}$ alkyl,
- 3) $(C=O)O_bC_3-C_8$ cycloalkyl,
- 4) $(C=O)O_b$ aryl,
- 5) $(C=O)O_b$ heterocyclyl,
- 6) C_1-C_{10} alkyl,
- 7) aryl,
- 8) C_2-C_{10} alkenyl,
- 9) C_2-C_{10} alkynyl,
- 10) heterocyclyl,
- 11) C_3-C_8 cycloalkyl,
- 12) SO_2R^a , and
- 13) $(C=O)NR^b_2$,

said alkyl, cycloalkyl, aryl, heterocyclyl, alkenyl, and alkynyl is optionally substituted with one, two or three substituents selected from R^7 , or

R^8 and R^9 can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R^7 ;

R^{10} is selected from: F and $-CH_2F$;

R^{13} is selected from: H and $-CH_2F$, provided that if t is 1, R^{13} is H;

R^{ox} is absent or is oxo;

R^a is independently selected from: (C_1-C_6) alkyl, (C_3-C_6) cycloalkyl, aryl, or heterocyclyl, optionally substituted with one, two or three substituents selected from R^7 ;

R^b is independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl, (C₃-C₆)cycloalkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'} or S(O)₂R^a, optionally substituted with one, two or three substituents selected from R⁷;

R^c and R^{c'} are independently selected from: H, (C₁-C₆)alkyl, aryl, NH₂, OH, OR^a, -(C₁-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'}, S(O)₂R^a and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷; or

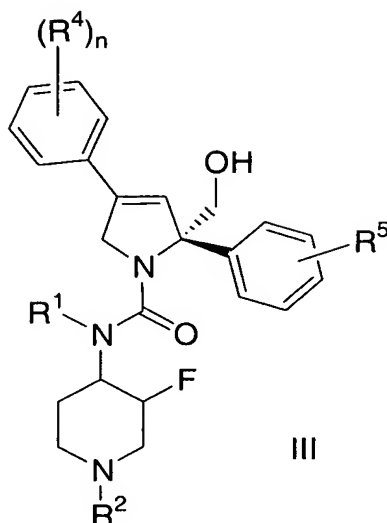
R^c and R^{c'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^d is selected from: H, (C₁-C₆)alkyl, -(C₂-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷;

R^e and R^{e'} are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷; or

R^e and R^{e'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷.

3. (Original) The compound according to Claim 2 of the Formula III:



or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

- a is 0 or 1;
b is 0 or 1;
m is 0, 1, or 2;
n is 0, 1 or 2;
r is 0 or 1;
s is 0 or 1;

R^1 and R^2 are independently selected from: H, (C₁-C₆)alkyl, aryl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R^7 ;

R^4 is independently selected from:

- 1) halo,
- 2) OH,
- 3) O_bC₁-C₆ perfluoroalkyl,

R^5 is selected from:

- 1) hydrogen,
- 2) halo,

- 3) OH,
- 4) $O_bC_1-C_6$ perfluoroalkyl,

R⁷ is selected from:

- 1) $(C=O)_rO_s(C_1-C_{10})$ alkyl,
- 2) $O_r(C_1-C_3)$ perfluoroalkyl,
- 3) oxo,
- 4) OH,
- 5) halo,
- 6) CN,
- 7) (C_2-C_{10}) alkenyl,
- 8) (C_2-C_{10}) alkynyl,
- 9) $(C=O)_rO_s(C_3-C_6)$ cycloalkyl,
- 10) $(C=O)_rO_s(C_0-C_6)$ alkylene-aryl,
- 11) $(C=O)_rO_s(C_0-C_6)$ alkylene-heterocyclyl,
- 12) $(C=O)_rO_s(C_0-C_6)$ alkylene-N(R^b)₂,
- 13) C(O)R^a,
- 14) (C_0-C_6) alkylene-CO₂R^a,
- 15) C(O)H,
- 16) (C_0-C_6) alkylene-CO₂H, and
- 17) C(O)N(R^b)₂,
- 18) S(O)_mR^a, and
- 19) S(O)₂N(R^b)₂;

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, alkylene and heterocyclyl is optionally substituted with up to three substituents selected from R^b, OH, (C₁-C₆)alkoxy, halogen, CO₂H, CN, O(C=O)C₁-C₆ alkyl, oxo, NO₂ and N(R^b)₂;

R⁸ and R⁹ are independently selected from:

- 1) H,
- 2) $(C=O)O_bC_1-C_{10}$ alkyl,
- 3) $(C=O)O_bC_3-C_8$ cycloalkyl,
- 4) $(C=O)O_b$ aryl,
- 5) $(C=O)O_b$ heterocyclyl,
- 6) C₁-C₁₀ alkyl,
- 7) aryl,

- 8) C₂-C₁₀ alkenyl,
- 9) C₂-C₁₀ alkynyl,
- 10) heterocyclyl,
- 11) C₃-C₈ cycloalkyl,
- 12) SO₂R^a, and
- 13) (C=O)NR^b₂,

said alkyl, cycloalkyl, aryl, heterocyclyl, alkenyl, and alkynyl is optionally substituted with one, two or three substituents selected from R⁷, or

R⁸ and R⁹ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^a is independently selected from: (C₁-C₆)alkyl, (C₃-C₆)cycloalkyl, aryl, or heterocyclyl, optionally substituted with one, two or three substituents selected from R⁷;

R^b is independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl, (C₃-C₆)cycloalkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'} or S(O)₂R^a, optionally substituted with one, two or three substituents selected from R⁷;

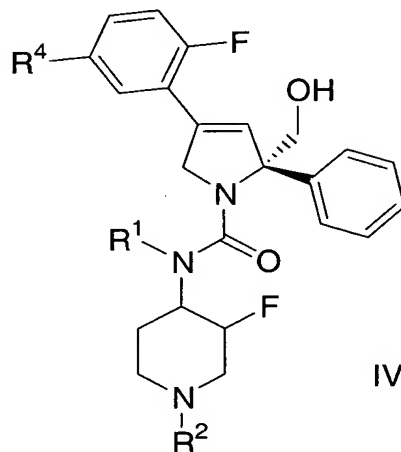
R^c and R^{c'} are independently selected from: H, (C₁-C₆)alkyl, aryl, NH₂, OH, OR^a, -(C₁-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'}, S(O)₂R^a and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷; or

R^c and R^{c'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^e and R^{e'} are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷; or

R^e and $R^{e'}$ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R^7 .

4. (Original) The compound according to Claim 3 of the Formula IV:



or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

- a is 0 or 1;
b is 0 or 1;
m is 0, 1, or 2;
r is 0 or 1;
s is 0 or 1;

R^1 and R^2 are independently selected from: H and (C_1-C_6) alkyl, optionally substituted with one, two or three substituents selected from R^7 ;

R^4 is independently selected from:

- 1) halo,
- 2) OH,
- 3) $O_bC_1-C_6$ perfluoroalkyl,

R⁷ is selected from:

- 1) (C=O)_rO_s(C₁-C₁₀)alkyl,
- 2) O_r(C₁-C₃)perfluoroalkyl,
- 3) oxo,
- 4) OH,
- 5) halo,
- 6) CN,
- 7) (C₂-C₁₀)alkenyl,
- 8) (C₂-C₁₀)alkynyl,
- 9) (C=O)_rO_s(C₃-C₆)cycloalkyl,
- 10) (C=O)_rO_s(C₀-C₆)alkylene-aryl,
- 11) (C=O)_rO_s(C₀-C₆)alkylene-heterocyclyl,
- 12) (C=O)_rO_s(C₀-C₆)alkylene-N(R^b)₂,
- 13) C(O)R^a,
- 14) (C₀-C₆)alkylene-CO₂R^a,
- 15) C(O)H,
- 16) (C₀-C₆)alkylene-CO₂H, and
- 17) C(O)N(R^b)₂,
- 18) S(O)_mR^a, and
- 19) S(O)₂N(R^b)₂;

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, alkylene and heterocyclyl is optionally substituted with up to three substituents selected from R^b, OH, (C₁-C₆)alkoxy, halogen, CO₂H, CN, O(C=O)C₁-C₆ alkyl, oxo, NO₂ and N(R^b)₂;

R⁸ and R⁹ are independently selected from:

- 1) H,
- 2) (C=O)O_bC₁-C₁₀ alkyl,
- 3) (C=O)O_bC₃-C₈ cycloalkyl,
- 4) (C=O)O_baryl,
- 5) (C=O)O_bheterocyclyl,
- 6) C₁-C₁₀ alkyl,
- 7) aryl,
- 8) C₂-C₁₀ alkenyl,
- 9) C₂-C₁₀ alkynyl,

- 10) heterocyclyl,
- 11) C₃-C₈ cycloalkyl,
- 12) SO₂R^a, and
- 13) (C=O)NR^b₂,

said alkyl, cycloalkyl, aryl, heterocyclyl, alkenyl, and alkynyl is optionally substituted with one, two or three substituents selected from R⁷, or

R⁸ and R⁹ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^a is independently selected from: (C₁-C₆)alkyl, (C₃-C₆)cycloalkyl, aryl, or heterocyclyl, optionally substituted with one, two or three substituents selected from R⁷;

R^b is independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl, (C₃-C₆)cycloalkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'} or S(O)₂R^a, optionally substituted with one, two or three substituents selected from R⁷;

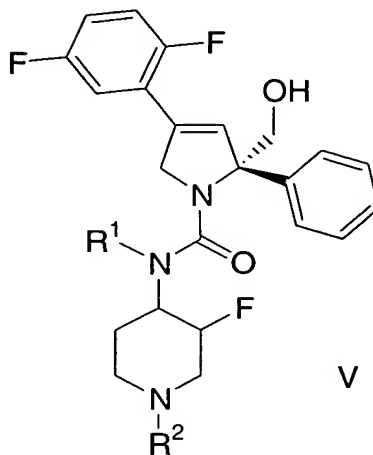
R^c and R^{c'} are independently selected from: H, (C₁-C₆)alkyl, aryl, NH₂, OH, OR^a, -(C₁-C₆)alkyl-OH, -(C₁-C₆)alkyl-O-(C₁-C₆)alkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl, (C=O)aryl, (C=O)heterocyclyl, (C=O)NR^eR^{e'}, S(O)₂R^a and -(C₁-C₆)alkyl-N(R^b)₂, wherein the alkyl is optionally substituted with one, two or three substituents selected from R⁷; or

R^c and R^{c'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷;

R^e and R^{e'} are independently selected from: H, (C₁-C₆)alkyl, aryl, heterocyclyl and (C₃-C₆)cycloalkyl, optionally substituted with one, two or three substituents selected from R⁷; or

R^e and R^{e'} can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 3-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one, two or three substituents selected from R⁷.

5. (Original) The compound according to Claim 4 of the Formula V:

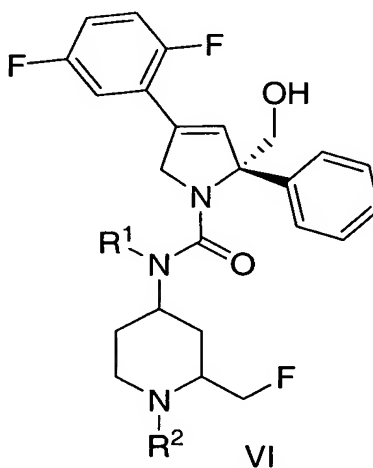


or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

R^1 and R^2 are independently selected from: H and (C_1-C_6) alkyl.

6. (Original) The compound according to Claim 2 of the formula VI:



or a pharmaceutically acceptable salt or stereoisomer thereof,

wherein:

R¹ and R² are independently selected from: H and (C₁-C₆)alkyl.

7. (Original) A compound selected from:

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*R*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoro-1-methylpiperidin-4-yl]-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*R*)-3-fluoro-1-methylpiperidin-4-yl]-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(2*R*,4*R*)-2-(fluoromethyl)-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(2*S*,4*S*)-2-(fluoromethyl)-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoro-1-methyl-1-oxidopiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

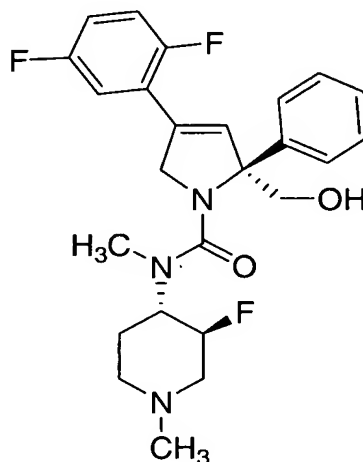
(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoropiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoro-1-isopropylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*S*)-3-fluoropiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

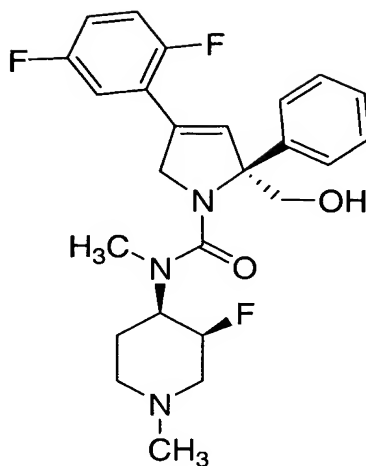
8. (Original) A compound which is:



(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

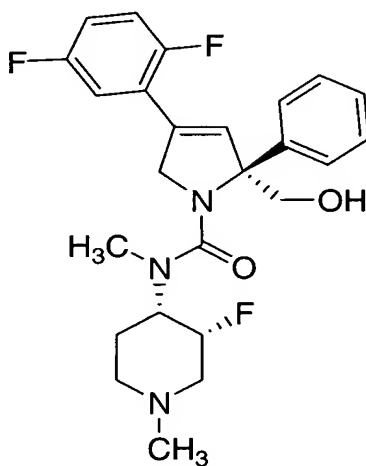
9. (Original) A compound which is:



(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*S*,4*R*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

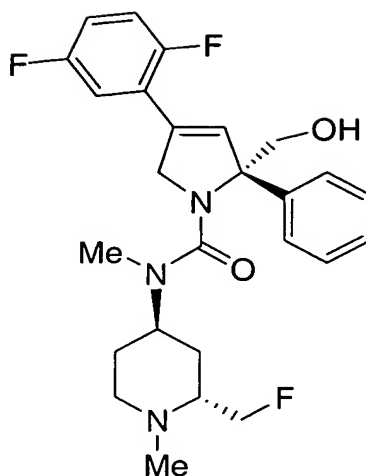
10. (Original) A compound which is:



(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

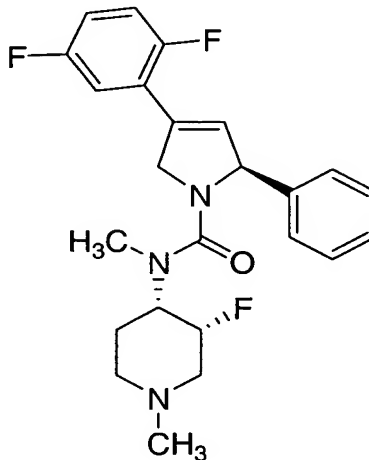
11. (Original) A compound which is:



(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(2*R*,4*R*)-2-(fluoromethyl)-1-methylpiperidin-4-yl]-2-(hydroxymethyl)-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

12. (Original) A compound which is:



(2*S*)-4-(2,5-Difluorophenyl)-*N*-[(3*R*,4*S*)-3-fluoro-1-methylpiperidin-4-yl]-*N*-methyl-2-phenyl-2,5-dihydro-1*H*-pyrrole-1-carboxamide

or a pharmaceutically acceptable salt thereof.

13. (Canceled)

14. (Original) A pharmaceutical composition that is comprised of a compound in accordance with Claim 1 and a pharmaceutically acceptable carrier.

15. (Original) A method of treating or preventing cancer in a mammal in need of such treatment that is comprised of administering to said mammal a therapeutically effective amount of a compound of Claim 1.

16. (Original) A method of treating cancer or preventing cancer in accordance with Claim 8 wherein the cancer is selected from cancers of the brain, genitourinary tract, lymphatic system, stomach, larynx and lung.

17. (Original) A method of treating or preventing cancer in accordance with Claim 8 wherein the cancer is selected from histiocytic lymphoma, lung adenocarcinoma, small cell lung cancers, pancreatic cancer, glioblastomas and breast carcinoma.

18. (Canceled)

19. (Canceled)

20. (Canceled)

21. (Canceled)

22. (Canceled)

23. (Canceled)

24. (Canceled)

25. (Canceled)

26. (Canceled)

27. (Original) A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy.

28. (Original) A method of treating or preventing cancer that comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a compound selected from:

- 1) an estrogen receptor modulator,
- 2) an androgen receptor modulator,
- 3) a retinoid receptor modulator,
- 4) a cytotoxic/cytostatic agent,
- 5) an antiproliferative agent,
- 6) a prenyl-protein transferase inhibitor,
- 7) an HMG-CoA reductase inhibitor,
- 8) an HIV protease inhibitor,
- 9) a reverse transcriptase inhibitor,
- 10) an angiogenesis inhibitor,
- 11) PPAR- γ agonists,
- 12) PPAR- δ agonists,
- 13) an inhibitor of inherent multidrug resistance,
- 14) an anti-emetic agent,
- 15) an agent useful in the treatment of anemia,
- 16) an agent useful in the treatment of neutropenia,
- 17) an immunologic-enhancing drug,
- 18) an inhibitor of cell proliferation and survival signaling, and
- 19) an agent that interferes with a cell cycle checkpoint.

29. (Original) A method of treating cancer that comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy and a compound selected from:

- 1) an estrogen receptor modulator,
- 2) an androgen receptor modulator,
- 3) a retinoid receptor modulator,
- 4) a cytotoxic/cytostatic agent,
- 5) an antiproliferative agent,

- 6) a prenyl-protein transferase inhibitor,
- 7) an HMG-CoA reductase inhibitor,
- 8) an HIV protease inhibitor,
- 9) a reverse transcriptase inhibitor,
- 10) an angiogenesis inhibitor,
- 11) PPAR- γ agonists,
- 12) PPAR- δ agonists,
- 13) an inhibitor of inherent multidrug resistance,
- 14) an anti-emetic agent,
- 15) an agent useful in the treatment of anemia,
- 16) an agent useful in the treatment of neutropenia,
- 17) an immunologic-enhancing drug,
- 18) an inhibitor of cell proliferation and survival signaling, and
- 19) an agent that interferes with a cell cycle checkpoint.

30. (Original) A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and paclitaxel or trastuzumab.

31. (Canceled)

32. (Canceled)

33. (Canceled)

34. (Original) A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a proteasome inhibitor.

35. (Original) A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with an aurora kinase inhibitor.

36. (Canceled)

37. (Original) A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a serine/threonine kinase inhibitor.

38. (Original) A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with an inhibitor of a mitotic kinesin that is not KSP.

39. (Original) A method of modulating mitotic spindle formation which comprises administering a therapeutically effective amount of a compound of Claim 1.

40. (Original) A method of inhibiting the mitotic kinesin KSP which comprises administering a therapeutically effective amount of a compound of Claim 1.